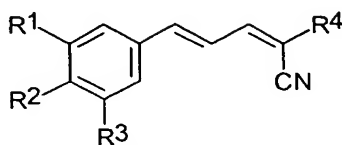


We claim:

1. A compound of Formula I, and salts, solvates or hydrates thereof:



5

wherein

- $R^1$  and  $R^2$  are each independently selected from the group consisting of H, OH,  $C_{1-6}$ alkyl,  $C_{1-6}$ alkoxy,  $NH_2$ ,  $NH-C_{1-6}$ alkyl,  $N(C_{1-6}alkyl)(C_{1-6}alkyl)$ , SH, S- $C_{1-6}$ alkyl, O-Si( $C_{1-6}alkyl$ )( $C_{1-6}alkyl$ )( $C_{1-6}alkyl$ ),  $NO_2$ ,  $CF_3$ ,  $OCF_3$  and halo;
- $R^3$  is selected from the group consisting of H, OH,  $C_{1-6}$ alkyl,  $C_{1-6}$ alkoxy,  $NH_2$ ,  $NH-C_{1-6}$ alkyl,  $N(C_{1-6}alkyl)(C_{1-6}alkyl)$ , SH, S- $C_{1-6}$ alkyl, O-Si( $C_{1-6}alkyl$ )( $C_{1-6}alkyl$ )( $C_{1-6}alkyl$ ),  $NO_2$ , halo and  $CH_2-S-(CH_2)_n$  Ar;
- $R^4$  is selected from the group consisting of  $C(X)R^5$ ,  $SO_3Ar$ ,  $NH_2$ ,  $NH-C_{1-6}alkyl$ ,  $N(C_{1-6}alkyl)(C_{1-6}alkyl)$ ,  $P(O)(OH)_2$ ,  $P(O)(OC_{1-6}alkyl)_2$ , and  $C(NH_2)=C(CN)_2$ ;
- X is selected from O, S, NH and N- $C_{1-6}alkyl$ ;
- $R^5$  is selected from the group consisting of  $NH_2$ , OH,  $NH(CH_2)_pAr$ ,  $NH(CH_2)_pOH$ ,  $(CH_2)_pOC_{1-6}alkyl$ ,  $C_{1-6}alkyl$ ,  $C_{1-6}alkoxy$ ,  $NHNH_2$ ,  $NHC(O)NH_2$ ,  $NHC(O)C_{1-6}alkoxy$ , N-morpholino and N-pyrrolidino; and
- Ar is an aromatic or heteroaromatic group, unsubstituted or substituted with 1-4 substituents, independently selected from the group consisting of OH,  $C_{1-6}alkyl$ ,  $C_{1-6}alkoxy$ ,  $NH_2$ ,  $NH-C_{1-6}alkyl$ ,  $N(C_{1-6}alkyl)(C_{1-6}alkyl)$ , SH, S- $C_{1-6}alkyl$ ,  $NO_2$ ,  $CF_3$ ,  $OCF_3$  and halo;
- n is 0 to 4; and
- p is 1-4.

25

2. The compound according to claim 1, wherein  $R^1$  and  $R^2$  are each independently selected from the group consisting of H, OH,  $C_{1-4}$ alkyl,  $C_{1-}$

alkoxy, NH<sub>2</sub>, NH-C<sub>1-4</sub>alkyl, SH, S-C<sub>1-4</sub>alkyl, O-Si(C<sub>1-4</sub>alkyl)(C<sub>1-4</sub>alkyl)(C<sub>1-4</sub>alkyl), NO<sub>2</sub>, CF<sub>3</sub>, OCF<sub>3</sub> and halo.

5        3.    The compound according to claim 2, wherein R<sup>1</sup> and R<sup>2</sup> are each independently selected from the group consisting of H, OH, OCH<sub>3</sub>, O-Si(CH<sub>3</sub>)<sub>2</sub><sup>t</sup>Bu, S-Me, SH and NO<sub>2</sub>.

10       4.    The compound according to claim 3, wherein R<sup>1</sup> and R<sup>2</sup> are both OH or R<sup>1</sup> and R<sup>2</sup> are both OCH<sub>3</sub>.

5       5.    The compound according to claim 4, wherein R<sup>1</sup> is OCH<sub>3</sub> and R<sup>2</sup> is OH.

15       6.    The compound according to claim 1, wherein R<sup>3</sup> is selected from the group consisting of H, OH, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkoxy, NH<sub>2</sub>, NH-C<sub>1-4</sub>alkyl, N(C<sub>1-4</sub>alkyl)(C<sub>1-4</sub>alkyl), SH, S-C<sub>1-4</sub>alkyl, NO<sub>2</sub> and halo.

7.    The compound according to claim 6, wherein R<sup>3</sup> is selected from the group consisting of H, OH, OCH<sub>3</sub>, SH, SMe, NO<sub>2</sub> and halo.

20       8.    The compound according to claim 7, wherein R<sup>3</sup> is selected from the group consisting of H, OH and OCH<sub>3</sub>.

25       9.    The compound according to claim 1, wherein R<sup>4</sup> is selected from the group consisting of C(X)R<sup>5</sup> and C(NH<sub>2</sub>)=C(CN)<sub>2</sub>.

10.   The compound according to claim 9, wherein R<sup>4</sup> is C(X)R<sup>5</sup>.

11.   The compound according to claim 10, wherein X is selected from the group consisting of O and S.

30       12.   The compound according to claim 10, wherein R<sup>5</sup> is selected from the group consisting of NH<sub>2</sub>, OH, NH(CH<sub>2</sub>)<sub>p</sub>Ar, NH(CH<sub>2</sub>)<sub>p</sub>OH and C<sub>1-4</sub>alkoxy.

13. The compound according to claim 12, wherein p is 1-3.
14. The compound according to claim 13, wherein R<sup>5</sup> is selected from the  
5 group consisting of NH<sub>2</sub>, OH, NH(CH<sub>2</sub>)<sub>p</sub>Ar, NH(CH<sub>2</sub>)<sub>p</sub>OH and OCH<sub>3</sub>.
15. The compound according to claim 14, wherein p is 1-2.
16. The compound according to claim 1, wherein Ar is an unsubstituted  
10 phenyl group or a phenyl group substituted with 1-4 substituents optionally  
selected from the group consisting of OH, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, NH<sub>2</sub>, NH-C<sub>1-6</sub>  
alkyl, N(C<sub>1-6</sub>alkyl)(C<sub>1-6</sub>alkyl), SH, S-C<sub>1-6</sub>alkyl, NO<sub>2</sub>, CF<sub>3</sub>, OCF<sub>3</sub> and halo.
17. The compound according to claim 14, wherein Ar is an unsubstituted  
15 phenyl group or a phenyl group substituted with 1-4 substituents optionally  
selected from the group consisting of OH, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, NH<sub>2</sub>, NH-C<sub>1-6</sub>  
alkyl, N(C<sub>1-6</sub>alkyl)(C<sub>1-6</sub>alkyl), SH, S-C<sub>1-6</sub>alkyl, NO<sub>2</sub>, CF<sub>3</sub>, OCF<sub>3</sub> and halo.
18. The compound according to any of claims 16 or 17, wherein Ar is an  
20 unsubstituted phenyl group or phenyl group substituted with 1-2 substituents  
optionally selected from the group consisting of OH, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkoxy,  
NH<sub>2</sub>, NH-C<sub>1-4</sub>alkyl, N(C<sub>1-4</sub>alkyl)(C<sub>1-4</sub>alkyl), SH, S-C<sub>1-4</sub>alkyl, NO<sub>2</sub>, CF<sub>3</sub>, OCF<sub>3</sub>  
and halo.
19. The compound according to claim 18, wherein Ar is an unsubstituted  
25 phenyl group or phenyl group substituted with 1-2 substituents optionally  
selected from the group consisting of OH, OCH<sub>3</sub>, NH<sub>2</sub>, NHCH<sub>3</sub>, N(CH<sub>3</sub>)<sub>2</sub>, SH,  
SCH<sub>3</sub>, CF<sub>3</sub>, OCF<sub>3</sub> and halo.
20. The compound according to claim 19, wherein Ar is selected from the  
30 group consisting of phenyl and 3,4-dihydroxyphenyl.

21. The compound according to claim 1, selected from the group consisting of:

(*E,E*)-2-(benzylamido)-3-styrylacrylonitrile (CR1);

(*E,E*)-2-(benzylamido)-3-(3,4-dimethoxystyryl)acrylonitrile (CR2);

5 (*E,E*)-2-(benzylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR3);

(*E,E*)-2-(benzylamido)-3-(3,4-dihydroxystyryl)acrylonitrile (CR4);

(*E,E*)-2-(phenylethylamido)-3-(3,4-dimethoxystyryl)acrylonitrile (CR5);

10 (*E,E*)-2-(phenylethylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR8);

(*E,E*)-2-(phenylpropylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR9);

(*E,E*)-2-(3,4-dihydroxybenzylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR11);

15 (*E,E*)-2-thioacetamido-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR12);

(*E,E*)-2-acetamido-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR13);

(*E,E*)-2-carboxy-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR14);

20 (*E,E*)-2-carbomethoxy-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR15);

(*E,E*)-2-acetamido-3-[3,4-bis(*t*-butyldimethylsilyloxystyryl)]acrylonitrile (CR16);

(*E,E*)-2-acetamido-3-(3,4-dihydroxystyryl)acrylonitrile (CR17);

25 (*E,E*)-2-(benzylamido)-3-(3,4-bis(*t*-butyldimethylsilyloxystyryl))acrylonitrile (CR18);

(*E,E*)-2-(3,4 dihydroxybenzylamido)-3-styrylacrylonitrile (CR19);

(*E,E*)-2-(3,4 dihydroxybenzylamido)-3-[3,4-bis(*t*-butyldimethylsilyloxystyryl)]acrylonitrile (CR20);

30 (*E,E*)-2-(3,4 dihydroxybenzylamido)-3-(3,4-dihydroxystyryl)acrylonitrile (CR21);

(*E,E*)-2-( $\beta$ -ethanolamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR24);

(*E,E*)-2-(benzylamido)-3-(4-nitrostyryl)acrylonitrile (CR27);

(*E,E*)-2-(3,4-dihydroxybenzylamido)-3-(4-nitrostyryl)acrylonitrile (CR28);

5 and

(*E,E*)-2-(1-amino-2,2-dicyanoethenyl)-3-(4-nitrostyryl)acrylonitrile (CR29).

22. The compound according to claim 21, selected from the group  
10 consisting of:

(*E,E*)-2-(benzylamido)-3-styrylacrylonitrile (CR1);

(*E,E*)-2-(benzylamido)-3-(3,4-dimethoxystyryl)acrylonitrile (CR2);

(*E,E*)-2-(benzylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR3);

15 (*E,E*)-2-(benzylamido)-3-(3,4-dihydroxystyryl)acrylonitrile (CR4);

(*E,E*)-2-(phenylethylamido)-3-(3,4-dimethoxystyryl)acrylonitrile (CR5);

(*E,E*)-2-(phenylpropylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR9);

20 (*E,E*)-2-(3,4-dihydroxybenzylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR11);

(*E,E*)-2-thioacetamido-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR12);

(*E,E*)-2-acetamido-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR13);

25 (*E,E*)-2-carboxy-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR14);

(*E,E*)-2-carbomethoxy-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR15);

(*E,E*)-2-acetamido-3-(3,4-dihydroxystyryl)acrylonitrile (CR17);

(*E,E*)-2-(3,4 dihydroxybenzylamido)-3-styrylacrylonitrile (CR19);

30 (*E,E*)-2-(3,4 dihydroxybenzylamido)-3-(3,4-dihydroxystyryl)acrylonitrile (CR21); and

(*E,E*)-2-( $\beta$ -ethanolamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR24).

23. The compound according to claim 22, selected from the group consisting of:

(*E,E*)-2-(benzylamido)-3-(3,4-dihydroxystyryl)acrylonitrile (CR4);

(*E,E*)-2-(3,4-dihydroxybenzylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR11);

(*E,E*)-2-acetamido-3-(3,4-dihydroxystyryl)acrylonitrile (CR17);

(*E,E*)-2-(3,4 dihydroxybenzylamido)-3-styrylacrylonitrile (CR19);

(*E,E*)-2-(3,4 dihydroxybenzylamido)-3-(3,4-dihydroxystyryl)acrylonitrile (CR21); and

(*E,E*)-2-( $\beta$ -ethanolamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR24).

24. The compound (*E,E*)-2-(benzylamido)-3-(3,4-dihydroxystyryl)acrylonitrile (CR4).

25. The compound (*E,E*)-2-(3,4-dihydroxybenzylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR11).

26. The compound (*E,E*)-2-(3,4-dihydroxybenzylamido)-3-(3,5-dimethoxy-4-hydroxystyryl)acrylonitrile (CR11).

27. A composition comprising a compound according to claim 1 in admixture with a pharmaceutically acceptable diluent or carrier.

28. A method of modulating cell proliferation comprising administering an effective amount of a compound of claim 23 to modulate cell proliferation to a cell or animal in need thereof.

29. A method of inhibiting cell proliferation comprising administering an effective amount of a compound of claim 23 to inhibit cell proliferation to a cell or animal in need thereof.

5 30. The method of claim 29, wherein the cell proliferation that is inhibited is cancer cell proliferation.

31. A method of treating cancer comprising administering to an animal in need thereof an effective amount of a compound of claim 23.

10

32. The method of claim 30 or 31 wherein said cancer is a hematopoietic cell cancer.

15 33. The method of claim 30 or 31 wherein said cancer is a leukemia, a lymphoma, a myeloma or a carcinoma.

20 34. The method of claim 33 wherein said leukemia is acute lymphoblastic leukemia, aggressive Philadelphia+ leukemia, acute myelocytic leukemia, chronic myeloid leukemia, chronic lymphocytic leukemia or juvenile myelomonocyte leukemia.

35. The method of claim 34 wherein said leukemia is acute lymphoblastic leukemia.

25 36. A method of modulating cell proliferation comprising administering an effective amount of a compound capable of modulating cell proliferation according to claim 1 or a composition of claim 27 to a cell or animal in need thereof.

30 37. A method of inhibiting cell proliferation comprising administering an effective amount of a compound capable of inhibiting cell proliferation

according to claim 1 or a composition according to claim 27 to a cell or animal in need thereof.

5        38.    A method of inhibiting cancer cell proliferation comprising administering an effective amount of a compound capable of inhibiting cancer cell proliferation according to any one of claim 1 or a composition according to claim 27 to a cell or animal in need thereof.

10       39.    A method of treating cancer comprising administering an effective amount of a compound capable of inhibiting cancer cell proliferation according to claim 1 or a composition according to claim 27 to a cell or animal in need thereof.

15       40.    A method according to claim 38 or 39 wherein said cancer is a hematopoietic cell cancer.

41.    A method according to claim 38 or 39 wherein said cancer is a leukemia, a lymphoma, a myeloma or a carcinoma.

20       41.    A method according to claim 41 wherein said leukemia is acute lymphoblastic leukemia, aggressive Philadelphia+ leukemia, acute myelocytic leukemia, chronic myeloid leukemia, chronic lymphocytic leukemia or juvenile myelomonocyte leukemia,

25       43.    A method according to claim 42 wherein said leukemia is acute lymphoblastic leukemia.